

**AMENDMENTS TO THE CLAIMS**

Please amend the claims to read as follows. Claims 22 and 53 are currently amended, with deleted text indicated by strikethrough, and claims 36, 49, 67, and 80 are canceled by this amendment.

- 1.-18. (canceled)
19. (previously presented) A method for treating a mycobacterial infection in a subject, the method comprising: administering to a subject an immunostimulatory nucleic acid molecule comprising an unmethylated CpG dinucleotide, in an amount effective to treat or ameliorate an infection with a *Mycobacterium* bacterium, thereby treating the infection in the subject.
20. (previously presented) The method of claim 19, wherein the immunostimulatory nucleic acid molecule is an immunostimulatory oligodeoxyribonucleotide.
21. (previously presented) The method of claim 19, wherein the immunostimulatory nucleic acid molecule is purified bacterial DNA.
22. (previously presented) The method of claim 19, wherein the immunostimulatory nucleic acid molecule is a plasmid DNA including sufficient immunostimulatory motifs to be immunostimulatory.
23. (previously presented) The method of claim 19, wherein the immunostimulatory nucleic acid molecule is a plasmid DNA which after being administered to the subject is degraded into oligonucleotides.
24. (withdrawn) The method of claim 19, wherein the immunostimulatory nucleic acid molecule comprises a CpG motif composed of an unmethylated CpG flanked by two 5'

purines and two 3' pyrimidines.

25. (previously presented) The method of claim 19, wherein the immunostimulatory nucleic acid molecule comprises a CpG motif in which the CpG is flanked by a 5' GpT dinucleotide and two 3' pyrimidines.

26. (previously presented) The method of claim 19, wherein the immunostimulatory nucleic acid molecule is 8-100 nucleotides long and comprises a CpG motif represented by:

$$5' X_1 X_2 C G X_3 X_4 3'$$

wherein C and G are unmethylated, and X<sub>1</sub>, X<sub>2</sub>, X<sub>3</sub> and X<sub>4</sub> are nucleotides.

27. (previously presented) The method of claim 19, wherein the immunostimulatory nucleic acid molecule is 8-100 nucleotides long and comprises a CpG motif represented by:

$$5' X_1 X_2 C G X_3 X_4 3'$$

wherein C and G are unmethylated, X<sub>1</sub>X<sub>2</sub> is selected from GpT, GpG, and GpA, and X<sub>3</sub> and X<sub>4</sub> are nucleotides.

28. (previously presented) The method of claim 19, wherein the immunostimulatory nucleic acid molecule is 8-100 nucleotides long and comprises a CpG motif represented by:

$$5' X_1 X_2 C G X_3 X_4 3'$$

wherein C and G are unmethylated, X<sub>1</sub> and X<sub>2</sub> are nucleotides, and X<sub>3</sub>X<sub>4</sub> is selected from TpT, CpT, and GpT.

29. (previously presented) The method of claim 19, wherein the immunostimulatory nucleic acid molecule is 8-100 nucleotides long and comprises a CpG motif represented by:

$$5' X_1 X_2 C G X_3 X_4 3'$$

wherein C and G are unmethylated, X<sub>1</sub>X<sub>2</sub> is selected from GpT, GpG, and GpA, and X<sub>3</sub>X<sub>4</sub> is selected from TpT, CpT, and GpT.

30. (previously presented) The method of claim 19, wherein the immunomodulatory nucleic acid molecule comprises a sequence selected from the group consisting of: AACGCC, AACGCT, AACGTC, AACGTT, AGCGCC, AGCGCT, AGCGTC, AGCGTT, GACGCC, GACGCT, GACGTC, GACGTT, GGCGCC, GGCGCT, GGCGTC, GGCGTT, ATCGCC, ATCGCT, ATCGTC, ATCGTT, GTCGCC, GTCGCT, GTCGTC, GTCGTT, and AACGCTCG.
31. (withdrawn) The method of claim 30, wherein the immunostimulatory nucleic acid molecule comprises the sequence AACGTT.
32. (previously presented) The method of claim 19, wherein said administering boosts the subject's immune response to eliminate an infection with a species of *Mycobacteria*.
33. (previously presented) The method of claim 19, wherein the bacterium is *Mycobacterium tuberculosis*.
34. (withdrawn) The method of claim 19, wherein the bacterium is *Mycobacterium avium*.
35. (previously presented) The method of claim 19, wherein the subject has an immune system deficiency.
36. (canceled)
37. (previously presented) A method for stimulating in a subject an immune response against a *Mycobacterium* bacterium, the method comprising: administering to a subject an immunostimulatory nucleic acid molecule comprising an unmethylated CpG dinucleotide, in an amount effective to stimulate an immune response in the subject, wherein said administering results in an immune response effective to treat, prevent or ameliorate an

infection in the subject, wherein the infection is a bacterial infection with a Mycobacterium bacterium.

38. (withdrawn) The method of claim 37, wherein the immunostimulatory nucleic acid molecule comprises a CpG motif composed of an unmethylated CpG flanked by two 5' purines and two 3' pyrimidines.
39. (previously presented) The method of claim 37, wherein the immunostimulatory nucleic acid molecule comprises a CpG motif in which the CpG is flanked by a 5' GpT dinucleotide and two 3' pyrimidines.
40. (previously presented) The method of claim 37, wherein the immunostimulatory nucleic acid molecule is 8-100 nucleotides long and comprises a CpG motif represented by:  
$$5' X_1 X_2 C G X_3 X_4 3'$$
wherein C and G are unmethylated, and X<sub>1</sub>, X<sub>2</sub>, X<sub>3</sub> and X<sub>4</sub> are nucleotides.
41. (previously presented) The method of claim 37, wherein the immunostimulatory nucleic acid molecule is 8-100 nucleotides long and comprises a CpG motif represented by:  
$$5' X_1 X_2 C G X_3 X_4 3'$$
wherein C and G are unmethylated, X<sub>1</sub>X<sub>2</sub> is selected from GpT, GpG, and GpA, and X<sub>3</sub> and X<sub>4</sub> are nucleotides.
42. (previously presented) The method of claim 37, wherein the immunostimulatory nucleic acid molecule is 8-100 nucleotides long and comprises a CpG motif represented by:  
$$5' X_1 X_2 C G X_3 X_4 3'$$
wherein C and G are unmethylated, X<sub>1</sub> and X<sub>2</sub> are nucleotides, and X<sub>3</sub>X<sub>4</sub> is selected from TpT, CpT, and GpT.
43. (previously presented) The method of claim 37, wherein the immunostimulatory

nucleic acid molecule is 8-100 nucleotides long and comprises a CpG motif represented by:



wherein C and G are unmethylated,  $X_1 X_2$  is selected from GpT, GpG, and GpA, and  $X_3 X_4$  is selected from TpT, CpT, and GpT.

44. (previously presented) The method of claim 37, wherein the immunostimulatory nucleic acid molecule comprises a sequence selected from the group consisting of: AACGCC, AACGCT, AACGTC, AACGTT, AGCGCC, AGCGCT, AGCGTC, AGCGTT, GACGCC, GACGCT, GACGTC, GACGTT, GGCGCC, GGCGCT, GGCGTC, GGCGTT, ATCGCC, ATCGCT, ATCGTC, ATCGTT, GTCGCC, GTCGCT, GTCGTC, GTCGTT, and AACGCTCG.
45. (withdrawn) The method of claim 44, wherein the immunostimulatory nucleic acid molecule comprises the sequence AACGTT.
46. (previously presented) The method of claim 37, wherein the bacterium is *Mycobacterium tuberculosis*.
47. (withdrawn) The method of claim 37, wherein the bacterium is *Mycobacterium avium*.
48. (previously presented) The method of claim 37, wherein the subject has an immune system deficiency.
49. (canceled)
50. (previously presented) A method for treating a mycobacterial infection in a subject, the method comprising: administering to a subject an immunostimulatory nucleic acid molecule comprising an unmethylated CpG dinucleotide, in an amount effective to treat,

prevent or ameliorate an infection in the subject, wherein the infection is a bacterial infection with a *Mycobacterium* bacterium, thereby treating the infection in the subject, wherein the immunostimulatory nucleic acid comprises an immunostimulatory sequence comprising an unmethylated cytosine, guanine dinucleotide sequence.

51. (previously presented) The method of claim 50, wherein the immunostimulatory nucleic acid molecule is an immunostimulatory oligodeoxyribonucleotide.
52. (previously presented) The method of claim 50, wherein the immunostimulatory nucleic acid molecule is purified bacterial DNA.
53. (previously presented) The method of claim 50, wherein the immunostimulatory nucleic acid molecule is a plasmid DNA including sufficient immunostimulatory motifs to be immunostimulatory.
54. (previously presented) The method of claim 50, wherein the immunostimulatory nucleic acid molecule is a plasmid DNA which after administration to the subject is degraded into oligonucleotides.
55. (withdrawn) The method of claim 50, wherein the immunostimulatory nucleic acid molecule comprises a CpG motif composed of an unmethylated CpG flanked by two 5' purines and two 3' pyrimidines.
56. (previously presented) The method of claim 50, wherein the immunostimulatory nucleic acid molecule comprises a CpG motif in which the CpG is flanked by a 5' GpT dinucleotide and two 3' pyrimidines.
57. (previously presented) The method of claim 50, wherein the immunostimulatory nucleic acid molecule is 8-100 nucleotides long and comprises a CpG motif represented by:

5' X<sub>1</sub>X<sub>2</sub>CGX<sub>3</sub>X<sub>4</sub> 3'

wherein C and G are unmethylated, and X<sub>1</sub>, X<sub>2</sub>, X<sub>3</sub> and X<sub>4</sub> are nucleotides.

58. (previously presented) The method of claim 50, wherein the immunostimulatory nucleic acid molecule is 8-100 nucleotides long and comprises a CpG motif represented by:

5' X<sub>1</sub>X<sub>2</sub>CGX<sub>3</sub>X<sub>4</sub> 3'

wherein C and G are unmethylated, X<sub>1</sub>X<sub>2</sub> is selected from GpT, GpG, and GpA, and X<sub>3</sub> and X<sub>4</sub> are nucleotides.

59. (previously presented) The method of claim 50, wherein the immunostimulatory nucleic acid molecule is 8-100 nucleotides long and comprises a CpG motif represented by:

5' X<sub>1</sub>X<sub>2</sub>CGX<sub>3</sub>X<sub>4</sub> 3'

wherein C and G are unmethylated, X<sub>1</sub> and X<sub>2</sub> are nucleotides, and X<sub>3</sub>X<sub>4</sub> is selected from TpT, CpT, and GpT.

60. (previously presented) The method of claim 50, wherein the immunostimulatory nucleic acid molecule is 8-100 nucleotides long and comprises a CpG motif represented by:

5' X<sub>1</sub>X<sub>2</sub>CGX<sub>3</sub>X<sub>4</sub> 3'

wherein C and G are unmethylated, X<sub>1</sub>X<sub>2</sub> is selected from GpT, GpG, and GpA, and X<sub>3</sub>X<sub>4</sub> is selected from TpT, CpT, and GpT.

61. (previously presented) The method of claim 50, wherein the immunostimulatory nucleic acid molecule comprises a sequence selected from the group consisting of: AACGCC, AACGCT, AACGTC, AACGTT, AGCGCC, AGCGCT, AGCGTC, AGCGTT, GACGCC, GACGCT, GACGTC, GACGTT, GGCGCC, GGCGCT, GGCGTC, GGCGTT, ATCGCC, ATCGCT, ATCGTC, ATCGTT, GTCGCC, GTCGCT, GTCGTC, GTCGTT, and AACGCTCG.

62. (withdrawn) The method of claim 61, wherein the immunostimulatory nucleic acid molecule comprises the sequence AACGTT.

63. (previously presented) The method of claim 50, wherein said administering boosts the subject's immune response to eliminate an infection with a species of *Mycobacteria*.

64. (previously presented) The method of claim 50, wherein the bacterium is *Mycobacterium tuberculosis*.

65. (withdrawn) The method of claim 50, wherein the bacterium is *Mycobacterium avium*.

66. (previously presented) The method of claim 50, wherein the subject has an immune system deficiency.

67. (canceled)

68. (previously presented) A method for stimulating in a subject an immune response against a *Mycobacterium* bacterium, the method comprising: administering to a subject an immunostimulatory nucleic acid molecule comprising an unmethylated CpG dinucleotide, in an amount effective to stimulate an immune response in the subject, wherein the immunostimulatory nucleic acid comprises an immunostimulatory sequence comprising an unmethylated cytosine, guanine dinucleotide sequence, wherein said administering results in an immune response effective to treat, prevent or ameliorate an infection in the subject, wherein the infection is a bacterial infection with a *Mycobacterium* bacterium.

69. (withdrawn) The method of claim 68, wherein the immunostimulatory nucleic acid molecule comprises a CpG motif composed of an unmethylated CpG flanked by two 5' purines and two 3' pyrimidines.

70. (previously presented) The method of claim 68, wherein the immunostimulatory nucleic acid molecule comprises a CpG motif in which the CpG is flanked by a 5' GpT dinucleotide and two 3' pyrimidines.

71. (previously presented) The method of claim 68, wherein the immunostimulatory nucleic acid molecule is 8-100 nucleotides long and comprises a CpG motif represented by:

$$5' X_1 X_2 C G X_3 X_4 3'$$

wherein C and G are unmethylated, and X<sub>1</sub>, X<sub>2</sub>, X<sub>3</sub> and X<sub>4</sub> are nucleotides.

72. (previously presented) The method of claim 68, wherein the immunostimulatory nucleic acid molecule is 8-100 nucleotides long and comprises a CpG motif represented by:

$$5' X_1 X_2 C G X_3 X_4 3'$$

wherein C and G are unmethylated, X<sub>1</sub>X<sub>2</sub> is selected from GpT, GpG, and GpA, and X<sub>3</sub> and X<sub>4</sub> are nucleotides.

73. (previously presented) The method of claim 68, wherein the immunostimulatory nucleic acid molecule is 8-100 nucleotides long and comprises a CpG motif represented by:

$$5' X_1 X_2 C G X_3 X_4 3'$$

wherein C and G are unmethylated, X<sub>1</sub> and X<sub>2</sub> are nucleotides, and X<sub>3</sub>X<sub>4</sub> is selected from TpT, CpT, and GpT.

74. (previously presented) The method of claim 68, wherein the immunostimulatory nucleic acid molecule is 8-100 nucleotides long and comprises a CpG motif represented by:

$$5' X_1 X_2 C G X_3 X_4 3'$$

wherein C and G are unmethylated, X<sub>1</sub>X<sub>2</sub> is selected from GpT, GpG, and GpA, and X<sub>3</sub>X<sub>4</sub> is selected from TpT, CpT, and GpT.

75. (previously presented) The method of claim 68, wherein the immunostimulatory nucleic acid molecule comprises a sequence selected from the group consisting of: AACGCC, AACGCT, AACGTC, AACGTT, AGCGCC, AGCGCT, AGCGTC, AGCGTT, GACGCC, GACGCT, GACGTC, GACGTT, GGCGCC, GGCGCT, GGCGTC, GGCGTT, ATCGCC, ATCGCT, ATCGTC, ATCGTT, GTCGCC, GTCGCT, GTCGTC, GTCGTT, and AACGCTCG.

76. (withdrawn) The method of claim 75, wherein the immunostimulatory nucleic acid molecule comprises the sequence AACGTT.

77. (previously presented) The method of claim 68, wherein the bacterium is *Mycobacterium tuberculosis*.

78. (withdrawn) The method of claim 68, wherein the bacterium is *Mycobacterium avium*.

79. (previously presented) The method of claim 68, wherein the subject has an immune system deficiency.

80. (canceled)

81. (previously presented) A method for treating a mycobacterial infection in a subject, the method comprising: administering to a subject an immunomodulatory nucleic acid molecule comprising an unmethylated CpG dinucleotide, in an amount effective to inhibit replication of a *Mycobacterium* bacterium, thereby treating mycobacterial infection in the subject.

82. (previously presented) The method of claim 81, wherein the immunomodulatory nucleic acid molecule is selected from the group consisting of an immunostimulatory

oligodeoxyribonucleotide (ISS-ODN); an isolated, detoxified bacterial polynucleotide; and an ISS-enriched plasmid DNA.

83. (previously presented) The method of claim 81, wherein the immunomodulatory nucleic acid molecule comprises a CpG motif selected from the group consisting of:
  - a) 5'-Purine-Purine-[C]-[G]-Pyrimidine-Pyrimidine-3';
  - b) 5'-Purine-TCG-Pyrimidine-Pyrimidine-3';
  - c) 5'-[TCG]<sub>n</sub>-3', where n is any integer that is at least 1; and
  - d) 5'-Purine-Purine-CG-Pyrimidine-Pyrimidine-CG-3'.
84. (previously presented) The method of claim 81, wherein the immunomodulatory nucleic acid molecule comprises a sequence selected from the group consisting of: AACGCC, AACGCT, AACGTC, AACGTT, AGCGCC, AGCGCT, AGCGTC, AGCGTT, GACGCC, GACGCT, GACGTC, GACGTT, GGCGCC, GGCGCT, GGCGTC, GGCGTT, ATCGCC, ATCGCT, ATCGTC, ATCGTT, GTCGCC, GTCGCT, GTCGTC, GTCGTT, and AACGCTCG.
85. (withdrawn) The method of claim 84, wherein the immunomodulatory nucleic acid molecule comprises the sequence AACGTT.
86. (previously presented) The method of claim 81, wherein said administering results in induction of an immune response effective against infection by a mycobacterial pathogen.
87. (previously presented) The method of claim 81, wherein the bacterium is *Mycobacterium tuberculosis*.
88. (withdrawn) The method of claim 81, wherein the bacterium is *Mycobacterium avium*.

89. (previously presented) The method of claim 81, wherein the subject is immunocompromised.

90. (previously presented) A method for inducing in a subject an immune response against a *Mycobacterium* bacterium, the method comprising: administering to a subject an amount of an immunomodulatory nucleic acid molecule comprising an unmethylated CpG dinucleotide, in an amount effective to elicit an immune response against a *Mycobacterium* bacterium; wherein said administering results in induction of an immune response effective to protect the subject against onset of disease or to decrease severity of symptoms of disease caused by infection by the *Mycobacterium* bacterium.

91. (previously presented) The method of claim 90, wherein the immunomodulatory nucleic acid molecule comprises a CpG motif selected from the group consisting of:

- 5'-Purine-Purine-[C]-[G]-Pyrimidine-Pyrimidine-3';
- 5'-Purine-TCG-Pyrimidine-Pyrimidine-3';
- 5'-[TCG]<sub>n</sub>-3', where n is any integer that is at least 1; and
- 5'-Purine-Purine-CG-Pyrimidine-Pyrimidine-CG-3'.

92. (previously presented) The method of claim 90, wherein the immunomodulatory nucleic acid molecule comprises a sequence selected from the group consisting of: AACGCC, AACGCT, AACGTC, AACGTT, AGCGCC, AGCGCT, AGCGTC, AGCGTT, GACGCC, GACGCT, GACGTC, GACGTT, GGGGCC, GGCGCT, GGCGTC, GGCGTT, ATCGCC, ATCGCT, ATCGTC, ATCGTT, GTGCGCC, GTGCGCT, GTGCGTC, GTGCGTT, and AACGCTCG.

93. (withdrawn) The method of claim 92, wherein the immunomodulatory nucleic acid molecule comprises the sequence AACGTT.

94. (previously presented) The method of claim 90, wherein the bacterium is *Mycobacterium tuberculosis*.

95. (withdrawn) The method of claim 90, wherein the bacterium is *Mycobacterium avium*.

96. (previously presented) The method of claim 90, wherein the subject is immunocompromised.

97. (previously presented) A method for treating a mycobacterial infection in a subject, the method comprising: administering to a subject multiple doses of an immunomodulatory nucleic acid molecule comprising an unmethylated CpG dinucleotide, in an amount effective to inhibit replication of a *Mycobacterium* bacterium, thereby treating the mycobacterial infection in the subject, wherein the immunomodulatory nucleic acid comprises an immunostimulatory sequence comprising 5' CpG 3'.

98. (previously presented) The method of claim 97, wherein the immunomodulatory nucleic acid molecule is selected from the group consisting of an immunostimulatory oligodeoxyribonucleotide (ISS-ODN); an isolated, detoxified bacterial polynucleotide; and an ISS-enriched plasmid DNA.

99. (previously presented) The method of claim 97, wherein the immunomodulatory nucleic acid molecule comprises a CpG motif selected from the group consisting of:  
5'-Purine-Purine-C-G-Pyrimidine-Pyrimidine-3';  
5'-Purine-TCG-Pyrimidine-Pyrimidine-3';  
5'-(TCG)<sub>n</sub>-3', where n is any integer that is at least 1; and  
5'-Purine-Purine-CG-Pyrimidine-Pyrimidine-CG-3'.

100. (previously presented) The method of claim 97, wherein the immunomodulatory nucleic acid molecule comprises a sequence selected from the group consisting of: AACGCC, AACGCT, AACGTC, AACGTT, AGCGCC, AGCGCT, AGCGTC, AGCGTT, GACGCC, GACGCT, GACGTC, GACGTT, GGCGCC, GGCGCT, GGCGTC, GGCGTT, ATCGCC, ATCGCT, ATCGTC, ATCGTT, GTCGCC, GTCGCT, GTCGTC, GTCGTT, and AACGCTCG.

101. (withdrawn) The method of claim 100, wherein the immunomodulatory nucleic acid molecule comprises the sequence AACGTT.

102. (previously presented) The method of claim 97, wherein said administering results in induction of an immune response effective against infection by a mycobacterial pathogen.

103. (previously presented) The method of claim 97, wherein the bacterium is *Mycobacterium tuberculosis*.

104. (withdrawn) The method of claim 97, wherein the bacterium is *Mycobacterium avium*.

105. (previously presented) The method of claim 97, wherein the subject is immunocompromised.

106. (previously presented) A method for inducing in a subject an immune response against a *Mycobacterium* bacterium, the method comprising: administering to a subject multiple doses of an immunomodulatory nucleic acid molecule comprising an unmethylated CpG dinucleotide, in an amount effective to elicit an immune response against a *Mycobacterium* bacterium, wherein the immunomodulatory nucleic acid comprises an immunostimulatory sequence comprising 5' CpG 3';

wherein said administering results in induction of an immune response effective to protect the subject against onset of disease or to decrease severity of symptoms of disease caused by infection by the *Mycobacterium* bacterium.

107. (previously presented) The method of claim 106, wherein the immunomodulatory nucleic acid molecule comprises a CpG motif selected from the group consisting of:

5'-Purine-Purine-C-G-Pyrimidine-Pyrimidine-3';  
5'-Purine-TCG-Pyrimidine-Pyrimidine-3';  
5'-(TCG)<sub>n</sub>-3', where n is any integer that is at least 1; and  
5'-Purine-Purine-CG-Pyrimidine-Pyrimidine-CG-3'.

108. (previously presented) The method of claim 106, wherein the immunomodulatory nucleic acid molecule comprises a sequence selected from the group consisting of:

AACGCC, AACGCT, AACGTC, AACGTT, AGCGCC, AGCGCT, AGCGTC, AGCGTT, GACGCC, GACGCT, GACGTC, GACGTT, GGCGCC, GGCGCT, GGCGTC, GGCGTT, ATCGCC, ATCGCT, ATCGTC, ATCGTT, GTCGCC, GTCGCT, GTCGTC, GTCGTT, and AACGCTCG.

109. (withdrawn) The method of claim 108, wherein the immunomodulatory nucleic acid molecule comprises the sequence AACGTT.

110. (previously presented) The method of claim 106, wherein the bacterium is *Mycobacterium tuberculosis*.

111. (withdrawn) The method of claim 106, wherein the bacterium is *Mycobacterium avium*.

112. (previously presented) The method of claim 106, wherein the subject is immunocompromised.